## **CLAIMS**

What is claimed is:

- 1. A polynucleotide vaccine comprising a nucleic acid sequence encoding an antigen derived from a non-host species of a first phylum or first kingdom, wherein the nucleic acid sequence encoding the antigen is modified by deletion of a native signal sequence.
- 2. The polynucleotide vaccine of claim 1, wherein the nucleic acid sequence encoding the antigen is further modified to include a signal sequence derived from a second phylum or second kingdom, wherein the signal sequence is operably linked to the antigen-encoding sequence.
- 3. The polynucleotide vaccine of claim 2, wherein the signal sequence comprises a hemagglutinin A (HA) signal sequence.
- 4. The polynucleotide vaccine of claim 1, wherein at least one codon of the nucleic acid sequence encoding the antigen is modified from a wild type sequence of the non-host species to an analogous codon of a host species.
- 5. The polynucleotide vaccine of claim 1, further comprising a universal antigen or an immunogenic fragment thereof.
  - 6. The polynucleotide vaccine of claim 1, wherein the first kingdom is plant.
  - 7. The polynucleotide vaccine of claim 1, wherein the antigen is Amb a1.
- 8. The polynucleotide vaccine of claim 1, wherein the antigen is derived from a pathogen.
- 9. The polynucleotide vaccine of claim 8, wherein the pathogen is a bacterium, a virus or a parasite.

- 10. A method for modulating an immune response to an antigen comprising administering to a subject a polynucleotide vaccine of any one of claims 1-9 in an amount effective to modulate an immune response to the antigen.
- 11. The method of claim 10, further comprising administering to the subject an immunostimulatory nucleotide sequence (ISS).
  - 12. The method of claim 10, wherein the antigen is an allergen.
- 13. The method of claim 12, wherein the allergen is a plant, food, latex, cat dander, cockroach or house dust mite allergen.
  - 14. The method of claim 13, wherein the plant allergen is ragweed or grass pollen.
- 15. A method for eliciting an immune response to an antigen comprising administering to a subject a polynucleotide vaccine of any one of claims 1-9 in an amount effective to elicit an immune response to the antigen.
- 16. The method of claim 15, further comprising administering to the subject an immunostimulatory nucleotide sequence (ISS).
  - 17. The method of claim 15, wherein the antigen is derived from a pathogen.
  - 18. The method of claim 17, wherein the pathogen is a bacterium, a virus or a parasite.
- 19. The method of claim 11, wherein the ISS comprises an unmethylated 5'-CG-3' nucleotide sequence.

- 20. The method of claim 19, wherein the ISS comprises a sequence selected from the group consisting of: 5'-rrcgyy-3', 5'-rrcgyy-3', 5'-rrcgyycg-3', 5'-rrcgyycg-3' or 5'-(TCG)n-3'.
- 21. The method of claim 20, wherein the sequence is selected from the group consisting of: AACGTT, AGCGTT, GACGTT, GGCGTT, AACGTC, AGCGTC, GACGTC, GGCGTC, AACGCC, AGCGCC, GACGCC, GGCGCC, AACGCT, AGCGCT, GACGCT, GGCGCT, ATCGTT, ACCGTT, GTCGTT, GCCGTT, ATCGTC, ACCGTC, GTCGTC, GCCGTC, ATCGCT, ACCGCT, GTCGCT, GCCGCT, ATCGCC, ACCGCC, GTCGCC, GCCGCC, AACGTTCG, AGCGTTCG, GACGTTCG, GACGTTCG, AACGTCCG, AACGTCCG, GACGTCCG, GACGCCCG, GACGCCCG, GACGCCCG, AACGTCCG, AACGTCCG, AACGCCCG, AACGCTCG, AACGCTCG, ACCGTCCG, ATCGTTCG, ACCGTTCG, GTCGTTCG, ATCGTTCG, ACCGTTCG, ATCGTTCG, GCCGTCCG, ATCGCCCG, ATCGCCCG, ACCGCCCG, ACCGCCCG, ATCGCCCG, ACCGCCCG, ACCGCCCG, ATCGCCCCG, ACCGCCCG, ACCGCCCG
- 22. A polynucleotide vaccine comprising a nucleic acid sequence encoding an Amb al allergen modified by deletion of a native Amb al signal sequence.
- 23. The polynucleotide vaccine of claim 22, wherein the nucleic acid sequence encoding the Amb a1 allergen is further modified to comprise a heterologous signal sequence operably linked to the Amb a1 allergen-encoding sequence.
- 24. The polynucleotide vaccine of claim 23, wherein the heterologous signal sequence comprises a hemagglutinin A (HA) signal sequence.
- 25. The polynucleotide vaccine of claim 22, wherein at least one codon of the nucleic acid sequence encoding the Amb a1 allergen is modified from a wild type sequence of the Amb a1 allergen to an analogous human codon.
  - 26. A polynucleotide vaccine composition comprising:

a polynucleotide comprising a nucleic acid sequence encoding an antigen derived from a non-host species of a first phylum or first kingdom, wherein the nucleic acid sequence encoding the antigen is modified by deletion of a native signal sequence; and

an immunomodulatory nucleic acid molecule comprising the sequence 5'-cytosine-guanine-3'.

- 27. The polynucleotide vaccine composition of claim 26, wherein the nucleic acid sequence encoding the antigen is further modified to include a heterologous signal sequence derived from a second phylum or second kingdom, wherein the signal sequence is operably linked to the antigen-encoding sequence.
- 28. The polynucleotide vaccine composition of claim 27, wherein the heterologous signal sequence comprises a hemagglutinin A (HA) signal sequence.
- 29. The polynucleotide vaccine composition of claim 26, wherein at least one codon of the nucleic acid sequence encoding the antigen is modified from a wild type sequence of the non-host species to an analogous codon of a host species.
- 30. The polynucleotide vaccine composition of claim 26, wherein the antigen is Amb a1.
- 31. The polynucleotide vaccine composition of claim 26, wherein the immunomodulatory nucleic acid molecule comprises a sequence selected from the group consisting of 5'-rrcgyy-3', 5'-rycgyy-3', 5'-rrcgyycg-3', 5'-rycgyycg-3' or 5'-(TCG)n-3'.
- 32. The polynucleotide vaccine composition of claim 26, wherein the immunomodulatory nucleic acid molecule comprises a sequence selected from the group consisting of: AACGTT, AGCGTT, GACGTT, GGCGTT, AACGTC, AGCGTC, GACGTC, GGCGTC, AACGCC, AGCGCC, GACGCC, GGCGCC, AACGCT, AGCGCT, GACGCT, GGCGCT, ATCGTT, ACCGTT, GTCGTT, GCCGTT, ATCGTC, ACCGTC, GTCGTC,